

GenCore version 5.1.6  
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OM protein - protein search, using swi model

Run on: June 25, 2003, 14:20:41 ; Search time 31.2 Seconds  
(without alignments)  
444.169 Million cell updates/sec

Title: US-09-622-613b-11  
Perfect score: 577  
Sequence: 1 SWMLPQKRLHFTNRDVCN.....TFCVTCENQAPVHFVGVC 104

Scoring table: BIOSUM62  
Gapop 10.0 , Gapext 0.5

Searched: 908470.seqs, 133250620 residues

Total number of hits satisfying chosen parameters: 908470

Minimum DB seq length: 0

Maximum DB seq length: 200000000

Post-processing: Minimum Match 0%  
Maximum Match 100%  
Listing first 45 summaries

Database :

1:	/SID52/gcgdata/geneseq/geneseq-emb1/AA1980.DAT.*
2:	/SID52/gcgdata/geneseq/geneseq-emb1/AA1981.DAT.*
3:	/SID52/gcgdata/geneseq/geneseq-emb1/AA1982.DAT.*
4:	/SID52/gcgdata/geneseq/geneseq-emb1/AA1983.DAT.*
5:	/SID52/gcgdata/geneseq/geneseq-emb1/AA1984.DAT.*
6:	/SID52/gcgdata/geneseq/geneseq-emb1/AA1985.DAT.*
7:	/SID52/gcgdata/geneseq/geneseq-emb1/AA1986.DAT.*
8:	/SID52/gcgdata/geneseq/geneseq-emb1/AA1987.DAT.*
9:	/SID52/gcgdata/geneseq/geneseq-emb1/AA1988.DAT.*
10:	/SID52/gcgdata/geneseq/geneseq-emb1/AA1989.DAT.*
11:	/SID52/gcgdata/geneseq/geneseq-emb1/AA1990.DAT.*
12:	/SID52/gcgdata/geneseq/geneseq-emb1/AA1991.DAT.*
13:	/SID52/gcgdata/geneseq/geneseq-emb1/AA1992.DAT.*
14:	/SID52/gcgdata/geneseq/geneseq-emb1/AA1993.DAT.*
15:	/SID52/gcgdata/geneseq/geneseq-emb1/AA1994.DAT.*
16:	/SID52/gcgdata/geneseq/geneseq-emb1/AA1995.DAT.*
17:	/SID52/gcgdata/geneseq/geneseq-emb1/AA1996.DAT.*
18:	/SID52/gcgdata/geneseq/geneseq-emb1/AA1997.DAT.*
19:	/SID52/gcgdata/geneseq/geneseq-emb1/AA1998.DAT.*
20:	/SID52/gcgdata/geneseq/geneseq-emb1/AA1999.DAT.*
21:	/SID52/gcgdata/geneseq/geneseq-emb1/AA2000.DAT.*
22:	/SID52/gcgdata/geneseq/geneseq-emb1/AA2001.DAT.*
23:	/SID52/gcgdata/geneseq/geneseq-emb1/AA2002.DAT.*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

## SUMMARIES

Result No.	Score	Query Match	Length DB	ID1	Description
1	577	100.0	104	AAV28870	Recombinant Rap1R1
2	577	100.0	105	AAV28871	Recombinant Met(-1
3	573	99.3	104	AAV28865	Rana pipiens liver
4	573	99.3	105	AAV28867	Recombinant Met(-1
5	573	99.3	127	AAV28879	Rana pipiens Clone
6	570	98.8	104	AAV28866	Recombinant Rap1R1
7	570	98.8	105	AAV28869	Recombinant Met(-1
8	555	96.4	104	AAW05544	Antitumor protein
9	555	96.2	112	AAW35116	R. pipiens recombi
10	555	96.2	251	AAW35134	R. pipiens recombi

11	555	96.2	254	18	AAW35135	R. pipiens recombi
12	555	96.2	355	18	AAW35129	R. pipiens recombi
13	555	96.2	355	18	AAW35133	R. pipiens recombi
14	555	96.2	366	18	AAW35132	R. pipiens recombi
15	551	95.5	104	12	AAV12344	Protein with activ
16	551	95.5	104	15	AAV47303	ONCONASR (pharmac
17	551	95.5	104	17	AAW0736	Protein derived fr
18	551	95.5	104	18	AAW30301	Recombinant onc pr
19	551	95.5	104	18	AAW06543	Antitumor protein
20	551	95.5	104	18	AAW14065	Onconase (RTM) pro
21	551	95.5	104	20	AAV33322	Frog onconase prot
22	551	95.5	104	20	AAW82233	Rana pipiens RNase
23	551	95.5	104	22	AAW31666	Amino acid sequenc
24	551	95.5	105	18	AAW35123	R. pipiens recombi
25	551	95.5	106	18	AAW35122	R. pipiens recombi
26	551	95.5	107	18	AAW35117	R. pipiens recombi
27	551	95.5	355	18	AAW35135	R. pipiens recombi
28	551	95.5	358	18	AAW35130	R. pipiens recombi
29	551	95.5	379	18	AAW35126	R. pipiens recombi
30	550	95.3	105	18	AAW35116	R. pipiens recombi
31	548	95.0	104	18	AAW30302	Recombinant onc pr
32	548	95.0	105	20	AAV39400	Recombinant frog O
33	546	94.6	104	18	AAW18224	Antitumor generic
34	546	94.6	105	18	AAW35115	R. pipiens recombi
35	546	94.6	358	18	AAW35127	R. pipiens recombi
36	546	94.6	365	18	AAW35131	R. pipiens recombi
37	543	94.1	104	22	AAW31667	Amino acid sequenc
38	531	92.0	107	18	AAW35120	R. pipiens recombi
39	494	85.6	360	18	AAW35128	R. pipiens recombi
40	484.5	84.0	111	18	AAW35121	R. pipiens recombi
41	445	77.1	83	20	AAW35119	R. pipiens clone R
42	445	77.1	83	20	AAW8234	Rana pipiens RNase
43	287	49.7	111	20	AAV33321	Frog lectin protol
44	280.5	48.6	110	20	AAV28877	Recombinant RacOR1
45	280.5	48.6	111	20	AAV28878	Recombinant Met(-1

## ALIGNMENTS

RESULT 1						
AAV28870						
ID	AAV28870 standard; Protein: 104 AA.					
XX						
AC	AAV28870;					
XX						
DT	25-JAN-2000 (first entry)					
XX						
DE	Recombinant Rap1R1 Gln1Ser amino acid sequence.					
XX						
KW	Recombinant Rana pipiens ribonuclease; Rap1R1 Gln1Ser; covalently bound;					
KW	IL2 antibody; ligand binding moiety; CD22; cancerous B cells; frog;					
KW	Kaposi's sarcoma; human chorionic gonadotropin; hCG; signal peptide;					
KW	recombinant ribonuclease; cytotoxic fusion protein; cancer; RNase;					
KW	autoimmune disease.					
XX						
OS	Rana pipiens.					
XX						
FT	Synthetic.					
FT						
FT	Key					
FT	Misc-difference 1					
FT	Location/Qualifiers					
FT	/note= "Wild type Gln replaced with Ser"					
PN	W09950398-AZ.					
XX						
PD	07-OCT-1999.					
XX						
PF	26-MAR-1999; 99WO-US06641.					
XX						
PR	27-MAR-1998; 98US-0079751.					
XX						
PA	(USSH ) US DEPT HEALTH & HUMAN SERVICES.					
XX						

PI Newton DL, Rybak SM;  
XX  
DR WPI: 1999-610847/52.  
DR N-PSDB: AA208128.  
XX  
PT New recombinant ribonucleases, used for killing target cells, e.g. for  
PT treating cancers, viral infections or autoimmune diseases  
XX  
PS Claim 34; Page 60; 71pp; English.  
XX  
CC The present sequence is a recombinant Rana pipiens ribonuclease (RaplR1)  
CC protein with Gln1ser. Carboxy terminal end of recombinant RaplR1 has a  
CC covalently bound ligand binding moiety, which can be a LL2 antibody  
CC directed against CD22 on cancerous B cells or human chorionic  
CC gonadotropin (hCG) effective against Kaposi's sarcoma cells. Recombinant  
CC ribonucleases can be expressed in bacteria without an N-terminal  
CC methionine due to the presence of a signal peptide that is cleaved by  
CC bacteria. The soluble expression of ribonuclease allows the proteins to  
CC be fused in-frame with ligand binding moieties to form cytotoxic fusion  
CC proteins. They can be used for treatment of cancer and autoimmune  
CC diseases.  
XX  
SQ Sequence 104 AA:  
XX  
Query Match 100.0%; Score 577; DB 20; Length 104;  
Best Local Similarity 100.0%; Pred. No. 2,9e-62;  
Matches 104; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
XX  
QY 1 SDMLTFQKKHLNTRDVCNNIMSTNLFHCCKDNFTYSRPPVKAICGIIASKNVLT 60  
DB 1 SDMLTFQKKHLNTRDVCNNIMSTNLFHCCKDNFTYSRPPVKAICGIIASKNVLT 60  
XX  
QY 61 SEFYISDCNVTSRPCKYKRLKSTNFTCVTCENQAPVHFVGHC 104  
DB 61 SEFYISDCNVTSRPCKYKRLKSTNFTCVTCENQAPVHFVGHC 104  
XX  
RESULT 2  
AA28871 ID AAY28871 standard; Protein: 105 AA.  
XX  
AC AAY28871;  
XX  
DT 25-JAN-2000 (first entry)  
XX  
DE Recombinant Met(-1) RaplR1 Gln1ser amino acid sequence.  
XX  
KW Recombinant Met(-1) Rana pipiens ribonuclease Gln1ser; RaplR1; CD22;  
KW covalently bound; LL2 antibody; ligand binding moiety; cancerous B cell;  
KW Kaposi's sarcoma; human chorionic gonadotropin; hCG; signal peptide;  
KW recombinant ribonuclease; cytotoxic fusion protein; cancer; frog;  
KW autoimmune disease; RNase.  
XX  
OS Rana pipiens.  
OS Synthetic.  
OS  
XX  
FH Key Location/Qualifiers  
FT Misc-difference 1 /note= "Met not found in wild type RaplR1"  
FT Misc-difference 2 /note= "Wild type Gln replaced with Ser"  
FT  
XX  
PN WO9950398-A2.  
XX  
PD 07-OCT-1999.  
XX  
PF 26-MAR-1999; 99WO-US06641.  
XX  
PR 27-MAR-1998; 98US-0079751.  
XX (USSH ) US DEPT HEALTH & HUMAN SERVICES.  
XX  
PI Newton DL, Rybak SM;

XX  
DR WPI: 1999-610847/52.  
DR N-PSDB: AA208129.  
XX  
PT New recombinant ribonucleases, used for killing target cells, e.g. for  
PT treating cancers, viral infections or autoimmune diseases  
XX  
PS Claim 34; Page 61; 71pp; English.  
XX  
CC The present sequence is a recombinant Rana pipiens ribonuclease (RaplR1)  
CC protein with Met at position 1 and Gln2Ser. Carboxy terminal end of  
CC recombinant RaplR1 has a covalently bound ligand binding moiety, which  
CC can be a LL2 antibody directed against CD22 on cancerous B cells or human  
CC chorionic gonadotropin (hCG) effective against Kaposi's sarcoma cells.  
CC Recombinant ribonucleases can be expressed in bacteria without an N-  
CC terminal methionine due to the presence of a signal peptide that is  
CC cleaved by bacteria. The soluble expression of ribonuclease allows the  
CC proteins to be fused in-frame with ligand binding moieties to form  
CC cytotoxic fusion proteins. They can be used for treatment of cancer and  
CC autoimmune diseases.  
XX  
SQ Sequence 105 AA:  
XX  
Query Match 100.0%; Score 577; DB 20; Length 105;  
Best Local Similarity 100.0%; Pred. No. 3e-62;  
Matches 104; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
XX  
QY 1 SDMLTFQKKHLNTRDVCNNIMSTNLFHCCKDNFTYSRPPVKAICGIIASKNVLT 60  
DB 2 SDMLTFQKKHLNTRDVCNNIMSTNLFHCCKDNFTYSRPPVKAICGIIASKNVLT 61  
XX  
QY 61 SEFYISDCNVTSRPCKYKRLKSTNFTCVTCENQAPVHFVGHC 104  
DB 62 SEFYISDCNVTSRPCKYKRLKSTNFTCVTCENQAPVHFVGHC 105  
XX  
RESULT 3  
AA28865 ID AAY28865 standard; Protein: 104 AA.  
XX  
AC AAY28865;  
XX  
DT 25-JAN-2000 (first entry)  
XX  
DE Rana pipiens liver ribonuclease (RaplR1).  
XX  
KW Rana pipiens liver ribonuclease; RaplR1; covalently bound; LL2 antibody;  
KW ligand-binding moiety; CD22; cancerous B cell; Kaposi's Sarcoma; frog;  
KW human chorionic gonadotropin; hCG; recombinant ribonuclease; RNase;  
KW signal peptide; cytotoxic fusion protein; cancer; autoimmune disease.  
XX  
OS Rana pipiens.  
OS  
XX  
PN WO9950398-A2.  
XX  
PD 07-OCT-1999.  
XX  
PF 26-MAR-1999; 99WO-US06641.  
XX  
PR 27-MAR-1998; 98US-0079751.  
XX (USSH ) US DEPT HEALTH & HUMAN SERVICES.  
XX  
PI Newton DL, Rybak SM;  
XX  
DR WPI: 1999-610847/52.  
DR N-PSDB: AA208124.  
XX  
PT New recombinant ribonucleases, used for killing target cells, e.g. for  
PT treating cancers, viral infections or autoimmune diseases  
XX  
PS Claim 1; Page 55; 71pp; English.

CC The present sequence is Rana pipiens liver ribonuclease (RaplR1)  
 CC protein. Carboxy terminal end of RaplR1 has a covalently bound  
 CC ligand binding moiety, which can be a LL2 antibody directed against  
 CC CD22 on cancerous B cells or human chorionic gonadotropin (hcg)  
 CC effective against Kaposi's Sarcoma cells. Recombinant ribonucleases can  
 CC be expressed in bacteria without an N-terminal methionine due to the  
 CC presence of a signal peptide that is cleaved by bacteria. The soluble  
 CC expression of ribonuclease allows the proteins to be fused in-frame with  
 CC ligand binding moieties to form cytotoxic fusion proteins. They can be  
 CC used for treatment of cancer and autoimmune diseases.

SO Sequence 104 AA:

Query Match

Best Local Similarity 100.0%; Score 573; DB 20; Length 104;

Matches 103; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 2 DMLTFQKKHLTNRDVDCNNIMSTNLFHCKDKNTFYISRPYPKAIKGIASKNVLTTS 61

DB 2 DMLTFQKKHLTNRDVDCNNIMSTNLFHCKDKNTFYISRPYPKAIKGIASKNVLTTS 61

OY 62 EFTLSDCNVTSRPPCKYKLRKSTNFCVTCENQAPVHFVGSHC 104

DB 62 EFTLSDCNVTSRPPCKYKLRKSTNFCVTCENQAPVHFVGSHC 104

RESULT 4

ID AAY28867 standard; Protein: 105 AA.

AC AAY28867;

DT 25-JAN-2000 (first entry)

XX Recombinant Met(-1) RaplR1.

DE Recombinant Met(-1) RaplR1.

XX Recombinant Met(-1) Rana pipiens ribonuclease; RaplR1; CD22; RNase;

KW covalently bound; LL2 antibody; ligand binding moiety; cancerous B cell;

KW Kaposi's sarcoma; human chorionic gonadotropin; hCG; signal peptide;

KW recombinant ribonuclease; cytotoxic fusion protein; cancer; frog;

XX autoimmune disease.

OS Rana pipiens.

OS Synthetic.

XX Key

FT Misc-difference 1

FT /note="Met not found in wild type RaplR1"

XX WO950398-A2.

XX 07-OCT-1999.

XX 26-MAR-1999; 99WO-US06641.

XX 27-MAR-1998; 98US-0079751.

XX (USSH ) US DEPT HEALTH & HUMAN SERVICES.

XX Newton DL, Rybak SM;

XX WPI: 1999-610847/52.

XX N-PSDB; AA208126.

XX New recombinant ribonucleases, used for killing target cells, e.g. for

XX treating cancers, viral infections or autoimmune diseases

CC gonadotropin (hCG) effective against Kaposi's sarcoma cells. Recombinant

CC ribonucleases can be expressed in bacteria without an N-terminal

CC methionine due to the presence of a signal peptide that is cleaved by

CC bacteria. The soluble expression of ribonuclease allows the proteins to

CC be fused in-frame with ligand binding moieties to form cytotoxic fusion

CC proteins. They can be used for treatment of cancer and autoimmune

CC diseases.

SO Sequence 105 AA:

Query Match

Best Local Similarity 100.0%; Score 573; DB 20; Length 105;

Matches 103; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 2 DMLTFQKKHLTNRDVDCNNIMSTNLFHCKDKNTFYISRPYPKAIKGIASKNVLTTS 61

DB 3 DMLTFQKKHLTNRDVDCNNIMSTNLFHCKDKNTFYISRPYPKAIKGIASKNVLTTS 62

OY 62 EFTLSDCNVTSRPPCKYKLRKSTNFCVTCENQAPVHFVGSHC 104

DB 63 EFTLSDCNVTSRPPCKYKLRKSTNFCVTCENQAPVHFVGSHC 105

RESULT 5

ID AAY28879 standard; Protein: 127 AA.

AC AAY28879;

DT 25-JAN-2000 (first entry)

XX Rana pipiens Clone 5a1b ribonuclease.

DE Rana pipiens ribonuclease Clone 5a1b; RaplR1; covalently bound; RNase;

KW LL2 antibody; ligand binding moiety; CD22; cancerous B cell; onconase;

KW Kaposi's Sarcoma; human chorionic gonadotropin; hCG; cancer;

KW recombinant ribonuclease; frog; signal peptide; cytotoxic fusion protein;

KW autoimmune disease.

XX Rana pipiens.

OS Key

FT Peptide

FT /label="Signal peptide"

FT /note="Putative"

FT Protein

FT /label="Rana\_pipiens\_Clone\_5a1b\_ribonuclease"

XX WO950398-A2.

XX 07-OCT-1999.

XX 26-MAR-1999; 99WO-US06641.

XX 27-MAR-1998; 98US-0079751.

XX (USSH ) US DEPT HEALTH & HUMAN SERVICES.

XX Newton DL, Rybak SM;

XX WPI: 1999-610847/52.

XX N-PSDB; AA208136.

XX New recombinant ribonucleases, used for killing target cells, e.g. for

XX treating cancers, viral infections or autoimmune diseases

XX Disclosure: Page 69; 71pp; English.

XX The present sequence is a Rana pipiens Clone 5a1b ribonuclease (RaplR1).

XX It is encoded by Clone 5a1b cDNA obtained from Rana pipiens liver mRNA

XX library. It exhibits differences with Onconase (RPM) at amino acid

XX residues 11, 20, 85 and 103. Carboxy terminal end of RaplR1 has a

XX covalently bound ligand binding moiety, which can be a LL2 antibody

CC directed against CD22 on cancerous B cells of human chorionic  
CC gonadotropin (hCG) effective against Kaposi's Sarcoma cells. Recombinant  
CC ribonucleases can be expressed in bacteria without an N-terminal  
CC methionine due to the presence of a signal peptide that is cleaved by  
CC bacteria. The soluble expression of ribonuclease allows the proteins to  
CC be fused in-frame with ligand binding moieties to form cytotoxic fusion  
CC proteins. They can be used for treatment of cancer and autoimmune  
CC diseases.

XX Sequence 127 AA;

Query Match 99.3%; Score 573; DB 20; Length 127;  
Best Local Similarity 100.0%; Pred. No. 1,2e-61;  
Matches 103; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 2 DWLTFQKKHLTNTRDVCNNIMSTNLFHCKDKNTFTYSRPEPKAICKGIASKNVLTTS 61  
DB 25 DWLTFQKKHLTNTRDVCNNIMSTNLFHCKDKNTFTYSRPEPKAICKGIASKNVLTTS 84  
QY 62 EFTLSDCNVTSRCKYKLLKSTNTFCVTCENQAPVHFVGSHC 104  
DB 85 EFTLSDCNVTSRCKYKLLKSTNTFCVTCENQAPVHFVGSHC 127

RESULT 6  
AAV28866  
ID AAV28866 standard; Protein: 104 AA.

XX AAV28866;

DT 25-JAN-2000 (first entry)

DE Recombinant RapL1 Met23Leu amino acid sequence.

XX Recombinant Rana pipiens ribonuclease: RapL1 Met23Leu; covalently bound;  
KW LL2 antibody; ligand binding moiety; CD22; cancerous B cell; RNase;  
KW Kaposi's sarcoma; human chorionic gonadotropin; hCG; signal peptide;  
KW recombinant ribonuclease; cytotoxic fusion protein; cancer; frog;  
KW autoimmune disease.

XX Rana pipiens.  
OS Synthetic.

XX Key Location/Qualifiers  
FT Misc-difference 23 /note= "Wild type Met replaced with Leu"

PN W09950398-A2.

PD 07-OCT-1999.

PF 26-MAR-1999; 99WO-US06641.

PR 27-MAR-1998; 98US-0079751.

XX (USSH ) US DEPT HEALTH & HUMAN SERVICES.

PA Newton DL, Rybak SM;

PI WPI: 1999-610847/52.

DR N-PSDB: AA208125.

PT New recombinant ribonucleases, used for killing target cells, e.g. for  
PT treating cancers, viral infections or autoimmune diseases

XX Claim 34; Page 56; 71pp: English.

XX The present sequence is a recombinant Rana pipiens ribonuclease (RapL1)  
CC protein with Met23Leu. Carboxy terminal end of recombinant RapL1 has a  
CC covalently bound ligand binding moiety, which can be a LL2 antibody  
CC directed against CD22 on cancerous B cells or human chorionic  
CC gonadotropin (hCG) effective against Kaposi's sarcoma cells. Recombinant  
CC ribonucleases can be expressed in bacteria without an N-terminal

CC methionine due to the presence of a signal peptide that is cleaved by  
CC bacteria. The soluble expression of ribonuclease allows the proteins to  
CC be fused in-frame with ligand binding moieties to form cytotoxic fusion  
CC proteins. They can be used for treatment of cancer and autoimmune  
CC diseases.

XX Sequence 104 AA;

Query Match 98.8%; Score 570; DB 20; Length 104;  
Best Local Similarity 99.0%; Pred. No. 2,1e-61;  
Matches 102; Conservative 1; Mismatches 0; Indels 0; Gaps 0;

QY 2 DWLTFQKKHLTNTRDVCNNIMSTNLFHCKDKNTFTYSRPEPKAICKGIASKNVLTTS 61  
DB 2 DWLTFQKKHLTNTRDVCNNIMSTNLFHCKDKNTFTYSRPEPKAICKGIASKNVLTTS 61  
QY 62 EFTLSDCNVTSRCKYKLLKSTNTFCVTCENQAPVHFVGSHC 104  
DB 62 EFTLSDCNVTSRCKYKLLKSTNTFCVTCENQAPVHFVGSHC 104

RESULT 7  
AAV28869  
ID AAV28869 standard; Protein: 105 AA.

XX AAV28869;

DT 25-JAN-2000 (first entry)

DE Recombinant Met(-1) RapL1 Met23Leu-(His)6 protein.

XX Recombinant Met(-1) Rana pipiens ribonuclease Met23Leu-(His)6: RapL1;  
KW CD22; covalently bound; LL2 antibody; ligand binding moiety; RNase;  
KW cancerous B cell; Kaposi's sarcoma; human chorionic gonadotropin; hCG;  
KW signal peptide; recombinant ribonuclease; cytotoxic fusion protein;  
KW cancer; frog; autoimmune disease.

XX Rana pipiens.  
OS Synthetic.

XX Key Location/Qualifiers

FT Misc-difference 1 /note= "(His)6 histidine tag attached to N-terminal Met"

FT Misc-difference 1 /note= "Met not found in wild type RapL1"

PN W09950398-A2.

PD 07-OCT-1999.

PF 26-MAR-1999; 99WO-US06641.

PR 27-MAR-1998; 98US-0079751.

XX (USSH ) US DEPT HEALTH & HUMAN SERVICES.

PA Newton DL, Rybak SM;

PI WPI: 1999-610847/52.

DR N-PSDB: AA208127.

PT New recombinant ribonucleases, used for killing target cells, e.g. for  
PT treating cancers, viral infections or autoimmune diseases

XX Claim 4; Page 59; 71pp: English.

XX The present sequence is a recombinant Rana pipiens ribonuclease protein  
CC (RapL1) with Met at position 1 attached to (His)6 tag and Met24Leu.  
CC Carboxy terminal end of recombinant RapL1 has a covalently bound  
CC binding moiety, which can be a LL2 antibody directed against CD22 on  
CC cancerous B cells or human chorionic gonadotropin (hCG) effective

CC against Kaposi's sarcoma cells. Recombinant ribonucleases can be  
 CC expressed in bacteria without an N-terminal methionine due to the  
 CC presence of a signal peptide that is cleaved by bacteria. The soluble  
 CC expression of ribonuclease allows the proteins to be fused in-frame with  
 CC ligand binding moieties to form cytotoxic fusion proteins. They can be  
 CC used for treatment of cancer and autoimmune diseases.

XX Sequence 105 AA:

Query Match 98.8%; Score 570; DB 20; Length 105;  
 Best Local Similarity 99.0%; Pred. No. 2, 1e-61;  
 Matches 102; Conservative 1; Mismatches 0; Indels 0; Gaps 0;

OY 2 DMLTFQKKHLNTRDVCNNIMSTNLFHCKDKNFTYSRPEPVKAICKGIIASKNVLTTS 61

DB 3 DMLTFQKKHLNTRDVCNNIMSTNLFHCKDKNFTYSRPEPVKAICKGIIASKNVLTTS 62

OY 62 EFYISDCNVTSRPCKYKILKSKSTNFCVCENQAPVHFVGVC 104

DB 63 EFYISDCNVTSRPCKYKILKSKSTNFCVCENQAPVHFVGVC 105

RESULT 8

AAW06544 ID AAW06544 standard; Protein; 104 AA.

XX AAW06544;

AC 22-AUG-1997 (first entry)

XX Antitumour protein from Rana pipiens oocytes.

DE Tumour; chemotherapy; radiotherapy; frog.

XX Rana pipiens.

XX WO9639428-A1.

PD 12-DEC-1996.

XX 03-JUN-1996; 96WO-US08304.

XX 06-JUN-1995; 95US-0467955.

XX (ALFA-) ALFACELL CORP.

XX Ardeli WT;

XX WPI; 1997-043063/C4.

XX Antitumour proteins from Rana pipiens oocytes - have fewer  
 CC disadvantages than chemotherapy, surgery and radiotherapy

XX Claim 8; Page 28; 45pp; English.

XX The present sequence is a specifically claimed example of an  
 CC antitumour protein from the generic protein in AAW8224, with the  
 CC molecular weight 12000. This is one of two preferred proteins (the  
 CC other in AAW06543) that have been isolated from Rana pipiens oocytes.  
 CC Both proteins have a blocked amino terminal group and are essentially  
 CC free of carbohydrates. The proteins are used to treat tumours. Use of  
 CC the peptides has fewer disadvantages than chemotherapy, radiotherapy  
 CC and surgery in the treatment of tumours.

XX Sequence 104 AA:

Query Match 96.4%; Score 556; DB 18; Length 104;  
 Best Local Similarity 97.1%; Pred. No. 1e-59;  
 Matches 100; Conservative 1; Mismatches 2; Indels 0; Gaps 0;

OY 2 DMLTFQKKHLNTRDVCNNIMSTNLFHCKDKNFTYSRPEPVKAICKGIIASKNVLTTS 61

DB 2 DMLTFQKKHLNTRDVCNNIMSTNLFHCKDKNFTYSRPEPVKAICKGIIASKNVLTTS 61

OY 62 EFYISDCNVTSRPCKYKILKSKSTNFCVCENQAPVHFVGVC 104

DB 62 EFYISDCNVTSRPCKYKILKSKSTNFCVCENQAPVHFVGVC 104

RESULT 9

AAW35118 ID AAW35118 standard; Protein; 112 AA.

XX AAW35118;

AC 20-APR-1998 (first entry)

XX R. pipiens recombinant RNase protein NLSmetseronc.

DE RNase A; ribonuclease; cytotoxic; onconase; none; immunofusion;

XX tumour cell growth; frog.

XX Rana pipiens.

XX WO9731116-A2.

XX 28-AUG-1997.

XX 19-FEB-1997; 97WO-US02588.

XX 21-FEB-1996; 96US-0011800.

XX (USSH ) US DEPT HEALTH & HUMAN SERVICES.

XX Boque L, Newton DL, Rybak SM, Wlodawer A;

XX WPI; 1997-435168/40.

XX N-PSDB; AAT94955.

XX Ribonuclease molecules based on native Onconase - used for killing  
 CC cells, particularly tumour cells

XX Claim 18; Page 63; 90pp; English.

XX AAW35115 to AAW35123 encode recombinant proteins (rOnc) which are  
 CC modifications of the RNase Onconase (rOnc). Such novel  
 CC ribonuclease molecules are highly cytotoxic and can be used alone or to  
 CC form chemical conjugates or to target recombinant immunofusions. They are  
 CC used particularly for decreasing tumour cell growth. They can also be  
 CC used for cell separation in vitro by selectively killing unwanted types  
 CC of cells, e.g. in bone marrow prior to transplantation into a patient  
 CC undergoing marrow ablation by radiation, or for killing leukemia cells.  
 CC or T-cells that would cause graft versus host disease. The toxins can  
 CC also be used to selectively kill unwanted cells in culture. The new  
 CC ribonucleases have increased cytotoxic activity compared to rOnc and also  
 CC lower immunogenicity in humans.

XX Sequence 112 AA:

Query Match 96.2%; Score 555; DB 18; Length 112;  
 Best Local Similarity 96.2%; Pred. No. 1.5e-59;  
 Matches 100; Conservative 2; Mismatches 2; Indels 0; Gaps 0;

OY 1 SOWLTFQKKHLNTRDVCNNIMSTNLFHCKDKNFTYSRPEPVKAICKGIIASKNVLTTS 60

DB 9 SOWLTFQKKHLNTRDVCNNIMSTNLFHCKDKNFTYSRPEPVKAICKGIIASKNVLTTS 68

OY 61 SEFYISDCNVTSRPCKYKILKSKSTNFCVCENQAPVHFVGVC 104

DB 69 SEFYISDCNVTSRPCKYKILKSKSTNFCVCENQAPVHFVGVC 112

RESULT 10

AAW35134 ID AAW35134 standard; Protein; 251 AA.

XX

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AC AAW35134;
XX
XX 20-APR-1998 (first entry)
XX
DE R. pipiens recombinant RNase ronc fusion protein 10.
XX
XX RNase A; ribonuclease; cytotoxic; onconase; nonc; immunofusion;
XX tumour cell growth; frog.
XX
OS Rana pipiens.
OS Synthetic.
XX
XX WO9731116-A2.
XX
XX 28-AUG-1997.
XX
XX 19-FEB-1997; 97WO-US02588.
XX
XX 21-FEB-1996; 96US-0011800.
XX
XX (USSH ) US DEPT HEALTH & HUMAN SERVICES.
XX
XX Boque L, Newton DL, Rybak SM, Wlodawer A;
XX
XX WPI: 1997-435168/40.
XX
XX N-PSDB: AAT94972.
XX
XX Ribonuclease molecules based on native Onconase - used for killing
XX cells, particularly tumour cells
XX
XX PS Disclosure: Page 76; 90pp; English.
XX
XX Sequences AAW35125 to AAW35135 represent recombinant fusion proteins
XX (ronc) which are modifications of the RNase Onconase (RTM) (nonc). Such
XX novel ribonuclease molecules are highly cytotoxic and can be used alone
XX or to form chemical conjugates or to target recombinant immunofusions.
XX They are used particularly for decreasing tumour cell growth. They can
XX also be used for cell separation in vitro by selectively killing unwanted
XX types of cells, e.g. in bone marrow prior to transplantation into a
XX patient undergoing marrow ablation by radiation, or for killing leukaemia
XX cells or T-cells that would cause graft versus host disease. The toxins
XX can also be used to selectively kill unwanted cells in culture. The new
XX ribonucleases have increased cytotoxic activity compared to nonc and
XX also lower immunogenicity in humans.
XX
XX SQ Sequence 251 AA;
XX
XX Query Match 96.2%; Score 555; DB 18; Length 251;
XX Best Local Similarity 96.2%; Pred. No. 4.3e-59;
XX Matches 100; Conservative 2; Mismatches 2; Indels 0; Gaps 0;
XX
XX QY 1 SDMLTFQKKHILNTROVDCNNIMSTNLFHCKDKNTFIYSRPEVKAICGIASKNVLT 60
XX DB 148 SDMLTFQKKHILNTROVDCNNIMSTNLFHCKDKNTFIYSRPEVKAICGIASKNVLT 207
XX
XX QY 61 SEFYLSDCNVTSRPCRYKLLKSTNTFCVCENQAPVHEVGVC 104
XX DB 208 SEFYLSDCNVTSRPCRYKLLKSTNTFCVCENQAPVHEVGVC 251
XX
XX RESULT 11
XX AAW35135
XX ID AAW35135 standard; Protein; 254 AA.
XX
XX AC AAW35135;
XX
XX XX 20-APR-1998 (first entry)
XX
XX DE R. pipiens recombinant RNase ronc fusion protein 11.
XX
XX XX RNase A; ribonuclease; cytotoxic; onconase; nonc; immunofusion;
XX tumour cell growth; frog.
XX
XX

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OS Rana pipiens.
OS Synthetic.
XX
XX WO9731116-A2.
XX
XX 28-AUG-1997.
XX
XX 19-FEB-1997; 97WO-US02588.
XX
XX 21-FEB-1996; 96US-0011800.
XX
XX (USSH ) US DEPT HEALTH & HUMAN SERVICES.
XX
XX Boque L, Newton DL, Rybak SM, Wlodawer A;
XX
XX WPI: 1997-435168/40.
XX
XX N-PSDB: AAT94973.
XX
XX Ribonuclease molecules based on native Onconase - used for killing
XX cells, particularly tumour cells
XX
XX PS Disclosure: Page 77; 90pp; English.
XX
XX Sequences AAW35125 to AAW35135 represent recombinant fusion proteins
XX (ronc) which are modifications of the RNase Onconase (RTM) (nonc). Such
XX novel ribonuclease molecules are highly cytotoxic and can be used alone
XX or to form chemical conjugates or to target recombinant immunofusions.
XX They are used particularly for decreasing tumour cell growth. They can
XX also be used for cell separation in vitro by selectively killing unwanted
XX types of cells, e.g. in bone marrow prior to transplantation into a
XX patient undergoing marrow ablation by radiation, or for killing leukaemia
XX cells or T-cells that would cause graft versus host disease. The toxins
XX can also be used to selectively kill unwanted cells in culture. The new
XX ribonucleases have increased cytotoxic activity compared to nonc and
XX also lower immunogenicity in humans.
XX
XX SQ Sequence 254 AA;
XX
XX Query Match 96.2%; Score 555; DB 18; Length 254;
XX Best Local Similarity 96.2%; Pred. No. 4.3e-59;
XX Matches 100; Conservative 2; Mismatches 2; Indels 0; Gaps 0;
XX
XX QY 1 SDMLTFQKKHILNTROVDCNNIMSTNLFHCKDKNTFIYSRPEVKAICGIASKNVLT 60
XX DB 2 SDMLTFQKKHILNTROVDCNNIMSTNLFHCKDKNTFIYSRPEVKAICGIASKNVLT 61
XX
XX QY 61 SEFYLSDCNVTSRPCRYKLLKSTNTFCVCENQAPVHEVGVC 104
XX DB 62 SEFYLSDCNVTSRPCRYKLLKSTNTFCVCENQAPVHEVGVC 105
XX
XX RESULT 12
XX AAW35129
XX ID AAW35129 standard; Protein; 355 AA.
XX
XX AC AAW35129;
XX
XX XX 20-APR-1998 (first entry)
XX
XX DE R. pipiens recombinant RNase ronc fusion protein 5.
XX
XX XX RNase A; ribonuclease; cytotoxic; onconase; nonc; immunofusion;
XX tumour cell growth; frog.
XX
XX OS Rana pipiens.
XX OS Synthetic.
XX
XX XX WO9731116-A2.
XX
XX XX 28-AUG-1997.
XX
XX XX 19-FEB-1997; 97WO-US02588.
XX
XX

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PR 21-FEB-1996; 9605-0011800.  
 XX (USSH ) US DEPT HEALTH & HUMAN SERVICES.  
 PA Bogue L, Newton DL, Rybak SM, Wlodawer A;  
 XX WPI; 1997-435168/40.  
 DR N-PSDB; AAT94971.  
 XX  
 PT Ribonuclease molecules based on native Onconase - used for killing  
 PT cells, particularly tumour cells;  
 PS Disclosure; Page 74; 90pp; English.  
 XX  
 CC Sequences AAW35125 to AAW35135 represent recombinant fusion proteins  
 CC (rOnc) which are modifications of the RNase Onconase (RM) (nOnc). Such  
 CC novel ribonuclease molecules are highly cytotoxic and can be used alone  
 CC or to form chemical conjugates or to target recombinant immunofusions.  
 CC They are used particularly for decreasing tumour cell growth. They can  
 CC also be used for cell separation in vitro by selectively killing unwanted  
 CC types of cells, e.g. in bone marrow prior to transplantation into a  
 CC patient undergoing marrow ablation by radiation, or for killing leukaemia  
 CC cells or T-cells that would cause graft versus host disease. The toxins  
 CC can also be used to selectively kill unwanted cells in culture. The new  
 CC ribonucleases have increased cytotoxic activity compared to nOnc and  
 CC also lower immunogenicity in humans.  
 CC  
 SQ Sequence 355 AA;  
 Query Match 96.2%; Score 555; DB 18; Length 355;  
 Best Local Similarity 96.2%; Pred. No. 6.7e-59;  
 Matches 100; Conservative 2; Mismatches 2; Indels 0; Gaps 0;  
 QY 1 SDMLTFQKKHLTNTRDVDCNNIMSTNLFHCKDKNTFIYSRPEPKAICKGIISKVLT 60  
 DB 252 SDMLTFQKKHLTNTRDVDCNNIMSTNLFHCKDKNTFIYSRPEPKAICKGIISKVLT 311  
 QY 61 SEFYLSDCNVTSRPCKYKLLKSTNTECVTCENQAPVHFVGVC 104  
 DB 312 SEFYLSDCNVTSRPCKYKLLKSTNTECVTCENQAPVHFVGVC 355  
 RESULT 13  
 AAW35133  
 ID AAW35133 standard; Protein: 355 AA.  
 AC AAW35133;  
 AC  
 AC  
 DT 20-APR-1998 (first entry)  
 DE R. pipiens recombinant RNase rOnc fusion protein 9.  
 XX  
 XX RNase A; ribonuclease; cytotoxic; onconase; nOnc; immunofusion;  
 KM tumour cell growth; frog.  
 XX  
 XX Rana pipiens.  
 OS Synthetic.  
 OS  
 PN WO9731116-A2.  
 PD 28-AUG-1997.  
 PD  
 PD 19-FEB-1997; 97WO-US02588.  
 PF 19-FEB-1997; 97WO-US02588.  
 PF  
 PR 21-FEB-1996; 96US-0011800.  
 PR  
 PA (USSH ) US DEPT HEALTH & HUMAN SERVICES.  
 PA Bogue L, Newton DL, Rybak SM, Wlodawer A;  
 PI WPI; 1997-435168/40.  
 DR N-PSDB; AAT94971.  
 XX

PT Ribonuclease molecules based on native Onconase - used for killing  
 PT cells, particularly tumour cells  
 XX  
 PS Disclosure; Page 75; 90pp; English.  
 XX  
 CC Sequences AAW35125 to AAW35135 represent recombinant fusion proteins  
 CC (rOnc) which are modifications of the RNase Onconase (RM) (nOnc). Such  
 CC novel ribonuclease molecules are highly cytotoxic and can be used alone  
 CC or to form chemical conjugates or to target recombinant immunofusions.  
 CC They are used particularly for decreasing tumour cell growth. They can  
 CC also be used for cell separation in vitro by selectively killing unwanted  
 CC types of cells, e.g. in bone marrow prior to transplantation into a  
 CC patient undergoing marrow ablation by radiation, or for killing leukaemia  
 CC cells or T-cells that would cause graft versus host disease. The toxins  
 CC can also be used to selectively kill unwanted cells in culture. The new  
 CC ribonucleases have increased cytotoxic activity compared to nOnc and  
 CC also lower immunogenicity in humans.  
 CC  
 SQ Sequence 355 AA;  
 Query Match 96.2%; Score 555; DB 18; Length 355;  
 Best Local Similarity 96.2%; Pred. No. 6.7e-59;  
 Matches 100; Conservative 2; Mismatches 2; Indels 0; Gaps 0;  
 QY 1 SDMLTFQKKHLTNTRDVDCNNIMSTNLFHCKDKNTFIYSRPEPKAICKGIISKVLT 60  
 DB 2 SDMLTFQKKHLTNTRDVDCNNIMSTNLFHCKDKNTFIYSRPEPKAICKGIISKVLT 61  
 QY 61 SEFYLSDCNVTSRPCKYKLLKSTNTECVTCENQAPVHFVGVC 104  
 DB 62 SEFYLSDCNVTSRPCKYKLLKSTNTECVTCENQAPVHFVGVC 105  
 RESULT 14  
 AAW35132  
 ID AAW35132 standard; Protein: 366 AA.  
 AC AAW35132;  
 AC  
 AC  
 DT 20-APR-1998 (first entry)  
 DE R. pipiens recombinant RNase rOnc fusion protein 8.  
 XX  
 XX RNase A; ribonuclease; cytotoxic; onconase; nOnc; immunofusion;  
 KM tumour cell growth; frog.  
 XX  
 XX Rana pipiens.  
 OS Synthetic.  
 OS  
 PN WO9731116-A2.  
 PD 28-AUG-1997.  
 PD  
 PD 19-FEB-1997; 97WO-US02588.  
 PF 19-FEB-1997; 97WO-US02588.  
 PF  
 PR 21-FEB-1996; 96US-0011800.  
 PR  
 PA (USSH ) US DEPT HEALTH & HUMAN SERVICES.  
 PA Bogue L, Newton DL, Rybak SM, Wlodawer A;  
 PI WPI; 1997-435168/40.  
 DR N-PSDB; AAT94970.  
 XX  
 XX Ribonuclease molecules based on native Onconase - used for killing  
 PT cells, particularly tumour cells  
 XX  
 PS Disclosure; Page 74; 90pp; English.  
 XX  
 CC Sequences AAW35125 to AAW35135 represent recombinant fusion proteins  
 CC (rOnc) which are modifications of the RNase Onconase (RM) (nOnc). Such  
 CC novel ribonuclease molecules are highly cytotoxic and can be used alone  
 CC or to form chemical conjugates or to target recombinant immunofusions.

CC They are used particularly for decreasing tumour cell growth. They can  
CC also be used for cell separation in vitro by selectively killing unwanted  
CC types of cells, e.g., in bone marrow prior to transplantation into a  
CC patient undergoing marrow ablation by radiation, or for killing leukaemia  
CC cells or T-cells that would cause graft versus host disease. The toxins  
CC can also be used to selectively kill unwanted cells in culture. The new  
CC ribonucleases have increased cytotoxic activity compared to none and  
CC also lower immunogenicity in humans.

XX Sequence 366 AA;

SO Query Match 96.2%; Score 555; DB 18; Length 366;  
Best Local Similarity 96.2%; Pred. No. 7e-59;

Matches 100; Conservative 2; Mismatches 2; Indels 0; Gaps 0;

QY 1 SDMLTFQKKHLTNRDVCNNIMSTNLFHCKDKNTFTYSRPEPVKAICKGIASKNVLTTS 60  
Db 263 SDMLTFQKKHLTNRDVCNNIMSTNLFHCKDKNTFTYSRPEPVKAICKGIASKNVLTTS 322

OY 61 SEFYLSDCNVTSRPCKYKTKKSTNFCVTCENQAPVHFVGSHC 104  
Db 323 SEFYLSDCNVTSRPCKYKTKKSTNFCVTCENQAPVHFVGSHC 366

RESULT 15

AA12344  
ID AA12344 standard; protein; 104 AA.

AC AA12344;

DT 08-AUG-1991 (first entry)

DE Protein with activity against cancer cells.

DE Frog eggs; Tamoxifen; Stelazine; cancer.

OS Rana pipiens.

PN W09107435-A.

PD 30-MAY-1991.

PF 26-OCT-1990; 90WO-US06185.

PR 18-MAY-1990; 90US-0526314.

PR 13-NOV-1989; 89US-0436141.

PA (ALFA-) ALFACELL CORP.

PI Ardelt WJ, Mikulski SM;

DR WPI; 1991-178059/24.

PT New protein from fertilised eggs of Rana pipiens - active against  
PT cancer cells, esp. in combination with Tamoxifen or Stelazine  
PT (trifluoro-per-azine).

PS Claim 7; Fig 2; 33pp; English.

XX The protein is derived from fertilised frog eggs. It has an iso-  
CC electric point of 9.5 - 10.5, a blocked N-terminal gp. and is free  
CC of carbohydrates. It is active against certain cancer cells. The  
CC combination of the protein and (2-1-p-dimethylaminoethoxyphenyl)-1,  
CC 2-diphenyl-1-butene) citrate salt (Tamoxifen) is much more bio-  
CC active than the separate entities against human pancreatic ASPC-1  
CC adenocarcinoma, and the combination of protein and (10-[3-(4-methyl  
CC piperazin-1-yl)-propyl]-2-trifluoromethylphenothiazine (Stelazine)  
CC is much more reactive than the separate entities against human lung  
CC A-549 carcinoma. Activity has also been shown against human sub-  
CC maxillary epidermoid carcinoma A-253 cells, human ovarian adeno-  
CC carcinoma NIH-OVCA3 cells, human leukaemic HL-60 cells, human  
CC COLO 320 DM cells, human LOX melanoma and human lung squamous car-  
CC cinoma HT-520 cells.

XX Sequence 104 AA;

SO Query Match 95.5%; Score 551; DB 12; Length 104;  
Best Local Similarity 96.1%; Pred. No. 4.2e-59;

Matches 99; Conservative 2; Mismatches 2; Indels 0; Gaps 0;

QY 2 DWLTFQKKHLTNRDVCNNIMSTNLFHCKDKNTFTYSRPEPVKAICKGIASKNVLTTS 61  
Db 2 DWLTFQKKHLTNRDVCNNIMSTNLFHCKDKNTFTYSRPEPVKAICKGIASKNVLTTS 61

OY 62 EFTLSDCNVTSRPCKYKTKKSTNFCVTCENQAPVHFVGSHC 104  
Db 62 EFTLSDCNVTSRPCKYKTKKSTNFCVTCENQAPVHFVGSHC 104

Search completed: June 25, 2003, 14:48:38  
Job time : 31.2 secs